FIG. 1A

Consensus Sequence

	1.00 1.00 1.00	50 DRSGNCVPCN DRSGNCVPCN DRSGNCVPCN DRSGNCVPCN	100 PCLDCAVVNR PCLDCAVVNR PCLDCAVVNR PCLDCAVVNR PCLDCAVVNR	150 PPPYEPHCA PPPYEPHCA PPPYEPHCA PPPYEPHCA
:: 6430	Weight: Weight: Weight:	SGDCRQQEFR 1 SGDCRQQEFR 1 TGDCRQQEFR 1 SGDCRQQEFR 1 SGDCRQQEFR 1	QCVTCRLHRF KEDWGFQKCK PCLDCAVVNR QCVTCRLHRF KEDWGFQKCK PCLDCAVVNR QCVTCRLHRF KEDWGFQKCK PCLDCAVVNR QCVACRLHRF KEDWGFQKCK PCLDCAVVNR QCVACRLHRF KEDWGFQKCK PCLDCAVVNR	
compCheck:	Check: 4050 We: Check: 4128 We: Check: 7937 We: Check: 8173 We:	GYLSCKVTCE GYLSCKVTCE GYLSCKVTCE GYLSCKVTCE	QCVTCRLHRF QCVTCRLHRF QCVTCRLHRF QCVACRLHRF	FYRKTKLVGF QDMECVPCGD FYRKTKLVGF QDMECVPCGD FYRKTKLVGF QDMECVPCGD FYRKTKLVGF QDMECVPCGD
blosum62.cmp	423 Chec) 423 Chec) 423 Chec) 423 Chec)	KTFFTLLVLL KTFFTLLVLL KTFFTLLVLL KTFFTLLVLL		
comparison table: 1	a 1 Len: Len: 1 Len: Len:	1 MALKVLLEQE MALKVLLEQE MALKVLLEQE MALKVLLEQE MALKVLLEQE	91 QCGPGMELSK ECGFGYGEDA QCGPGMELSK ECGFGYGEDA QCGPGMELSK ECGFGYGEDA QCGPGMELSK ECGFGYGEDA	FOKANCSATS DAICGDCLPG FOKANCSATS DAICGDCLPG FOKANCSATS DAICGDCLPG FOKANCSATS DAICGDCLPG FOKANCSATS DAICGDCLPG
ol compariso		hTRADEalpha_ hTRAIN_1 hTRADEbeta_1 hApo420pep Consensus	hTRADEalpha_ hTRAIN_1 hTRADEbeta_1 hApo420pep Consensus	hTRADEalpha_ hTRAIN_1 hTRADEbeta_1 hApo420pep Consensus
Symbol	Name: Name: Name:	htrad htrad htrad hap Co	htrad h htrad hap	hTRAD h' hTRADI hAP

フォ

FIG. 1B

	151				200
hTRADEalpha	SKVNLVKIAS	TASSPRDTAL	AAVICSALAT	SKVNLVKIAS TASSPRDTAL AAVICSALAT VLLALLILCV IYCKRQFMEK	IYCKRQFMEK
hTRAIN_1	SKVNLVKIAS	TASSPRDTAL	AAVICSALAT	TASSPRDTAL AAVICSALAT VLLALLILCV	IYCKROFMEK
hTRADEbeta_1	SKVNLVKIAS		AAVICSALAT	TASSPRDTAL AAVICSALAT VLLALLILCV	IYCKRQFMEK
hApo420pep	SKVNLVKIAS	TASSPRDTAL	AAVICSALAT	SKVNLVKIAS TASSPRDTAL AAVICSALAT VLLALLILCV IYCKRQFMEK	IYCKRQFMEK
Consensus	SKVNLVKIAS	TASSPRDTAL	AAVICSALAT	SKVNLVKIAS TASSPRDTAL AAVICSALAT VLLALLILCV IYCKRQFMEK	IYCKRQFMEK
	201				C
hTRADEalpha	KPSWSLRSQD	IQYNGSELSC	FDRPOLHEYA	KPSWSLRSOD IQYNGSELSC FDRPOLHEYA HRACCOCRRD SVOTCGPVRL	SVOTCGPVRL
hTRAIN 1	KPSWSLRSQD	IQYNGSELSC	FDRPQLHEYA	IQYNGSELSC FDRPQLHEYA HRACCQCRRD	SVOTCGPVRL
hTRADEbeta_1	KPSWSLRSQD		LDRPQLHEYA	IQYNGSELSC LDRPQLHEYA HRACCQCRRD SVQTCGPVRL	SVOTCGPVRL
hApo420pep	KPSWSLRSQD	IQYNETELSC	FDRPQLHEYA	KPSWSLRSQD IQYNETELSC FDRPQLHEYA HRACCQCRRD SVQTCGPVRL	SVQTCGPVRL
Consensus	KPSWSLRSQD	IQYNGSELSC	FDRPQLHEYA	KPSWSLRSQD IQYNGSELSC FDRPQLHEYA HRACCQCRRD SVQTCGPVRL	SVQTCGPVRL
	251				300
hTRADEalpha_	LPSMCCEEAC	SPNPATLGCG	VHSAASLQAR	LPSMCCEEAC SPNPATLGCG VHSAASLQAR NAGPAGEMVP TFFGSLTQSI	TFFGSLTQSI
hTRAIN_1	LPSMCCEEAC	SPNPATLGCG	VHSAASLQAR	LPSMCCEEAC SPNPATLGCG VHSAASLQAR NAGPAGEMVP TFFGSLTQSI	TFFGSLTQSI
					t

Constant Con

hTRADEbeta_1 hApo420pep Consensus

LPSMCCEEAC SPNPATLGCG VHSAASLQAR NAGPAGEMVP TFFGSLTQSI

FIG. 1C

350	rsedsn Fsedsn	LESSTSIDSN	ISLDSN	ISLDSN	400	RSQLDQ	RSQLDQ	RSQLDQ	RSQLDQ	RSQLDQ					
	NPE LESS	NPE LESS	NPE LESS	NPE LESS		TOD ALTM									
	GEDIHSL	GEDIHSL	GEDIHSL	GEDIHSL		TLVESAS	TLVESAS	TLVESAS	TLVESAS	TLVESAS					
	SFCDSYPELT GEDIHSLNPE LESSTSLDSN		SFCDSYPELT GEDIHSLNPE LESSTSLDSN	SPCDSYPELT		AATDLSRYNN	AATDLSRYNN	AATDLSRYNN	AATDLSRYNN	AATDLSRYNN	423	1 1	1	GSL	ממד.
	CGEFSDAWPL MONPMGGDNI SFCDSYPELT GEDIHSLNPE LESSTSLDSN	CGEFSDAWFL MONFMGGDNI	CGEFSDAWPL MONPMGGDNI	CGEFSDAWPL MONPMGGDNI SPCDSYPELT GEDIHSLNPE LESSTSLDSN		SSQDLVGGAV PVQSHSENFT AATDLSRYNN TLVESASTQD ALTMRSQLDQ	SSQDLVGGAV PVQSHSENFT AATDLSRYNN TLVESASTQD ALTMRSQLDQ	SSQDLVGGAV PVQSHSENFT AATDLSRYNN TLVESASTQD ALTMRSQLDQ	SSODLVGGAV PVOSHSENFT AATDLSRYNN TLVESASTOD ALTMRSQLDQ	SSQDLVGGAV PVQSHSENFT AATDLSRYNN TLVESASTQD ALTMRSQLDQ		ESGALIHPAT QTSLQEA~~~	ESGAVIHPAT QTSLQEA~~~	ESGAIIHPAT QTSLQVRQRL GSL	TOUR THE OWEL OVER DE
301	CGEFSDAWPL	CGEFSDAWFL	CGEFSDAWPL	CGEFSDAWPL	351	SSQDLVGGAV	SSQDLVGGAV	SSQDLVGGAV	SSQDLVGGAV	SSQDLVGGAV	401	ESGALIHPAT	ESGAVIHPAT	ESGALIHPAT	日本のは上げるこのは
	hTRADEalpha	hTRADEbeta 1	hApo420pep	Consensus		hTRADEalpha	hTRAIN 1	hTRADEbeta 1	hApo420pep	Consensus		hTRADEalpha	hTRAIN 1	hTRADEbeta_1	L 2 m C L C L C L

ESGAVIHPAT QTSLQVRQRL GSL ESGA-IHPAT QTSLQ--QRL GSL

hApo420pep Consensus

FIG. 2

TRADE- α ESGAIIHPATQTSLQEA TRADE- β ESGAIIHPATQTSLQVRQRLGSL 401 416 423

FIG. 3

CRD1

NGFRp75	CPTGLYT.H	SGEC CKAC	NLGEGVAQPC	G.ANQTVCE
OX40				SRSQNTVCR
CD40				TEFTETECL
TRADE	CRQQEFRDR	SGNCVPCNQC	GPGMELSKEC	GFGYGEDAQCV

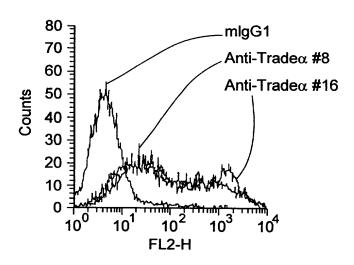
CRD2

NGFRp75				MSAPCVE	
OX40	PCGPGFY	NDVVSSKPCK	PCTWC.NLRS	GSERKQLCTA	TQDTVC
CD40	PCGES.EFL	DTWNRETHCH	QHKYCDPNLG	LRVQQKGTSE	T.DTIC
TRADE	TCRLHR.FK	EDWGFQK.CK	PCLDCAVVN.	.rfqkancsa	TSDAIC

FIG. 4

50	100	150 150	200	250 250	300	350 349	400 399	
→ MALKVILEQEKTFFILLVILGYLSCKVICESGI MALKVIPLHRTVLEAAILFLHLACKVSCEFGI	CDR2 QCGPGMELSKECGFGYGEDAQCVTCRLHRFKEDWGFQKCKPCLDCAVVNR QCGPGMELSKECGFGYGEDAQCVPCRPHRFKEDWGFQKCKPCADCALVNR	FQKANCSALSDALCGDCLPGFYRKTKLVGFQDMECVPCGDPPPPYEPHCA FQKANCSALSDAVCGDCLPGFYRKTKLVGFQDMECVPCGDPPPPYEPHCL	SKVNLVKIASTASSPRDTALAAVICSALATVLLALLILCVIYCKRQFMEK SKVNLVKISSTASSPRDTALAAVICSALATVLLALLILCVIYCKRQFMEK	KPSWSLRSQDIQYNGSELSCFDRPDLHEYAHRACCOCRRDSVQTCGPVRL KPSWSLRSQDIQYNGSELSCFDQPRLRHCAHRACCQYHRDSAPMYGPVHL	LPSMCCEEACSPNPATLGCCVHSAASLOARNAGPAGEMVPTFFGSLTOSI IPSLCCEEARSSARAVLGCGLRSPTTLOERNPASVGDTMPAFFGSVSRSI	CGEFSDAWPLMONPMGGDNISFCDSYPELTGEDIHSLNPELESSTSLDSN CAEFSDAWPLMONPLGGDS-SLCDSYPELTGEDINSLNPENESAASLDSS	SSDDLWGGAVPVQSHSENFTAATDLSRYNNTLVESASTQDALTMRSQLDQ GGQDLAGTAALESSGNVSESTDSPRHGDTGTVWEQTJAQDAQRTPSQGGW	ESGAVIHPATOTSLOER 417 EDRENLNLAMPTAFODA 416
SEQ ID NO:2HUMAN TRADE $lpha$ SEQ ID NO:6MOUSE TRADE	SEQ ID NO:2HUMAN TRADE $lpha$ SEQ ID NO:6MOUSE TRADE	SEQ ID NO:2HUMAN TRADE $lpha$ SEQ ID NO:6MOUSE TRADE	SEQ ID NO:2HUMAN TRADE α SEQ ID NO:6MOUSE TRADE	SEQ ID NO:2HUMAN TRADE $lpha$ SEQ ID NO:6MOUSE TRADE	SEQ ID NO:2HUMAN TRADE $lpha$ SEQ ID NO:6MOUSE TRADE	SEQ ID NO:2HUMAN TRADE α SEQ ID NO:6MOUSE TRADE	SEQ ID NO:2HUMAN TRADE $lpha$ SEQ ID NO:6MOUSE TRADE	SEQ ID NO: ZHUMAN TRADE $lpha$ SEQ ID NO: 6MOUSE TRADE

FIG. 5A



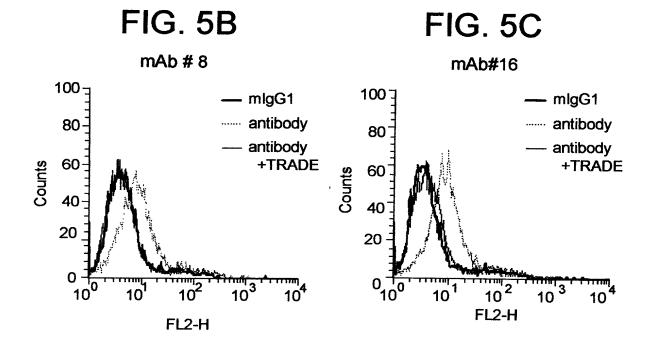


FIG. 6A

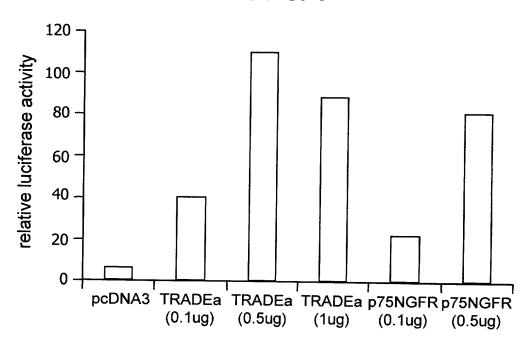
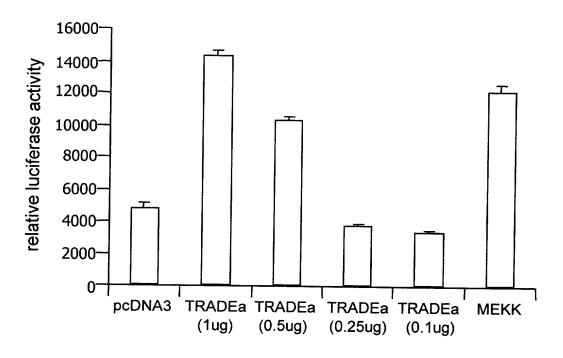
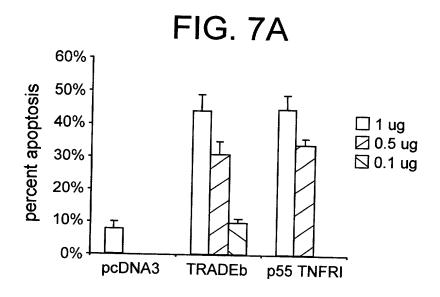
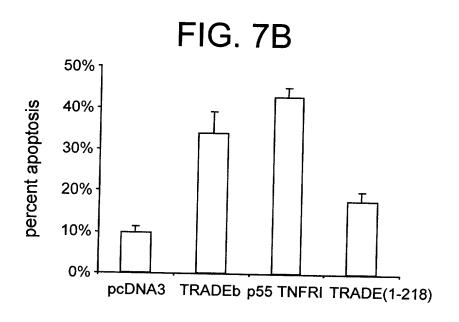


FIG. 6B







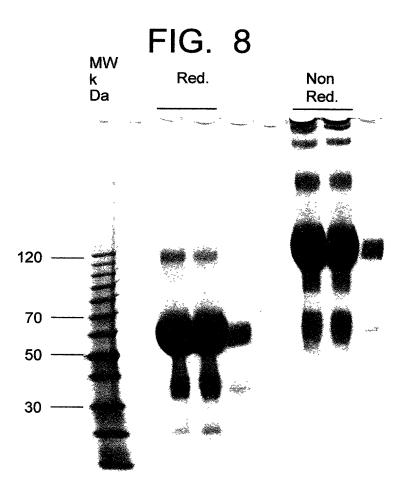
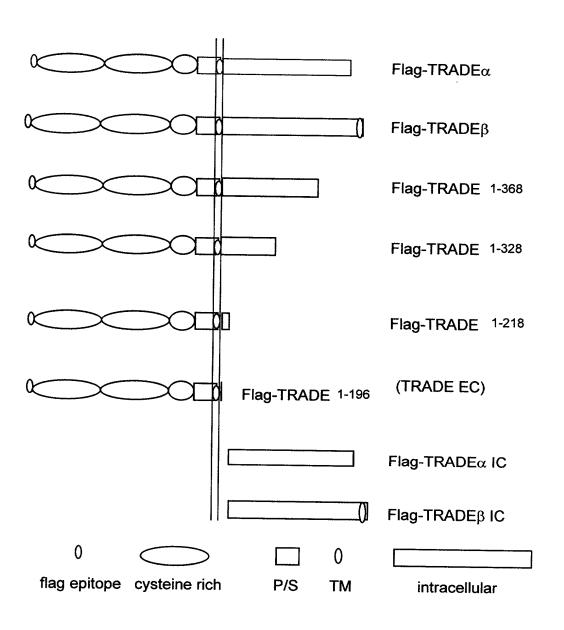


FIG. 9



Deletion analysis

FIG. 10

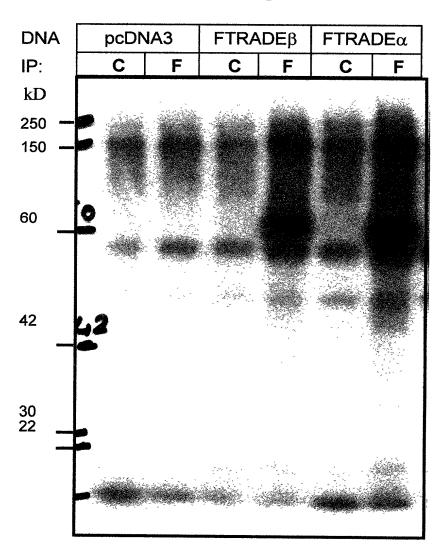
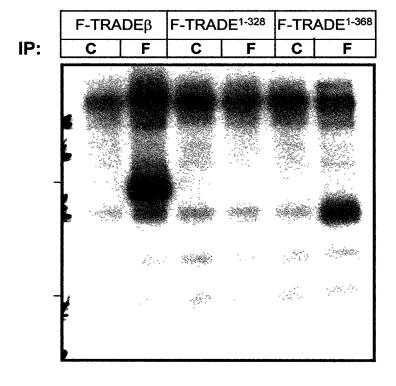
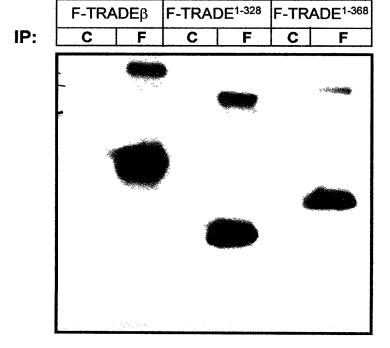


FIG. 11A



Kinase assay

FIG. 11B



anti-Flag blot

FIG. 12A

 $HA-TRAF6(\Delta N) +$

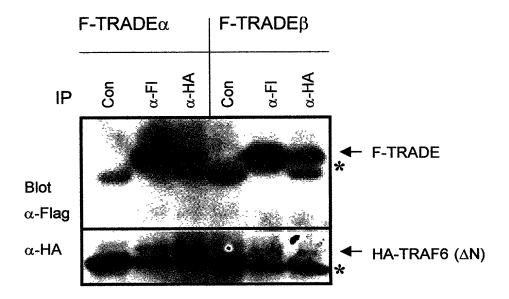


FIG. 12B



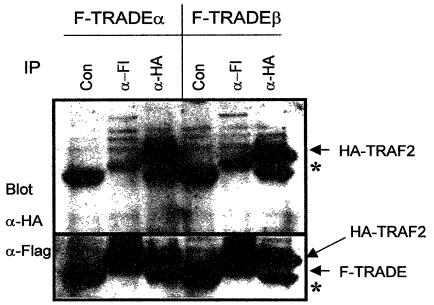


FIG. 13

HA-TRAF3+

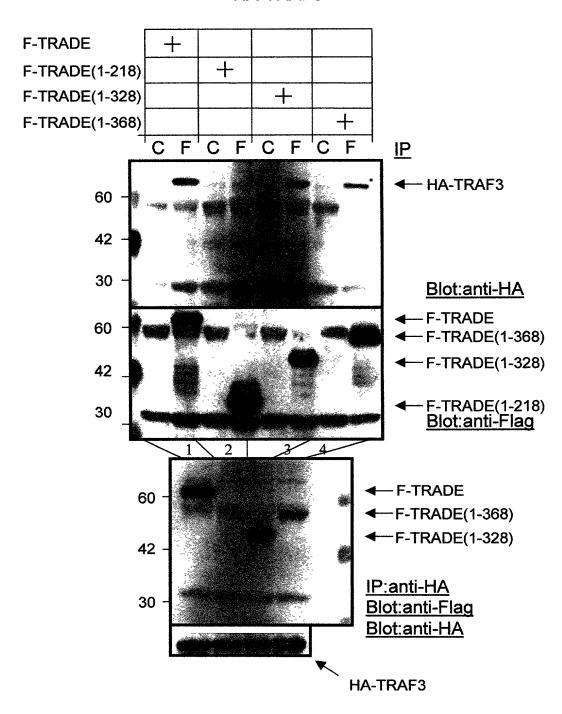


FIG. 14A

